

CLAIMS

What is claimed is:

- 5 1. A vector system for transfection and recombinant polypeptide expression in a mammalian host cell comprising:
 - (a) a first cistron encoding a transactivator protein under control of a first promoter;
and
 - (b) a second cistron encoding an apoptosis-protective protein under the control of
10 the first promoter or optionally under the control of a second promoter;
wherein the first and the second cistron are contained in one or more vectors.
2. The vector system of Claim 1, further comprising a third cistron encoding at least one desired polypeptide under control of a third promoter, wherein said third promoter is responsive to the transactivator protein and wherein the first, the second, and the third
15 cistrons are contained in one or more vectors.
3. The vector system of Claim 2, further comprising one or more additional cistrons each encoding a desired polypeptide under control of a promoter responsive to the transactivator protein.
4. The vector system of Claim 2, wherein said polypeptide is a single chain antibody or
20 a heavy or light chain of an antibody or antibody fragment.
5. The vector system of Claim 1, wherein the first and second cistrons are on one vector and the first cistron lies downstream of the second cistron.
6. The vector system of Claim 1, wherein the first cistron encodes a CREB protein or a variant thereof.
- 25 7. The vector system of Claim 6, wherein the CREB protein variant is CREB variant Y134F.
8. The vector system of Claim 6, wherein the second cistron encodes an adenoviral E1b-19K protein, a Bcl-2 protein, or a Bcl-2 protein having a deletion in the regulatory loop domain.

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20. The vector system of Claim 19, wherein the ubiquitous chromatin opening element comprises an extended methylation-free CpG-island.

21. The vector system of Claim 19, wherein the ubiquitous chromatin opening element comprises a hnRNP A2 promoter.

5 22. A method of expressing a desired recombinant polypeptide in a mammalian host cell comprising introducing to the mammalian host cell:

(a) a first cistron encoding a transactivator protein under control of a first promoter;

(b) a second cistron encoding an apoptosis-protective protein under control of the first promoter or optionally under control of a second promoter; and

10 (c) a third cistron encoding the desired polypeptide under control of a third promoter;

wherein said third promoter is responsive to the transactivator-protein. .

23. The method of Claim 22, wherein the third cistron is associated with a ubiquitous chromatin opening element, an insulator, or a barrier element.

15 24. The method of Claim 22, wherein the host cell is selected from the group consisting of a CHO cell, a mouse myeloma cell, a mouse hybridoma cell, a rat myeloma cell, and a rat hybridoma cell.

25. The method of Claim 24, wherein the host cell is a cell capable of growing in a suspension.

20 26. The method of Claim 24, wherein the host cell is a YB2/0 rat hybridoma cell.

27. The method of Claim 22, wherein the first or second promoter is an efficient heterologous promoter.

28. The method of Claim 22, wherein the transactivator and the apoptotic protective protein are homologous to the endogenous transactivator and apoptotic protective proteins
25 of the host cell.

29. The method of Claim 22, wherein the first cistron encodes a transactivator protein selected from the group consisting of an E1a protein, a CREB protein, and variants thereof.

9. The vector system of Claim 1, wherein the first cistron encodes an adenoviral E1a polypeptide or a variant thereof.
10. The vector system of Claim 9, wherein the adenoviral E1a variant comprises a mutation in CR1.
- 5 11. The vector system of Claim 10, wherein the adenoviral E1a variant comprises a 47H mutation.
12. The vector system of Claim 1, wherein the second cistron encodes an apoptosis-protective protein selected from the group consisting of a dominant negative mutant of p53, a protein that interacts with BAX, a protein that interacts with BAK, an inhibitor of
10 apoptosome formation, and a downstream apoptosis inhibitor.
13. The vector system of Claim 1, wherein the second cistron encodes an adenoviral E1b-19K protein, a Bcl-2 protein, or a Bcl-2 protein having a deletion in the regulatory loop domain.
14. The vector system of Claim 1, wherein the first or second promoter is an efficient
15 heterologous promoter.
15. The vector system of Claim 1, wherein the first or second promoter is a RSV-LTR promoter, a SV-40 promoter, or a cytomegalovirus promoter.
16. The vector system of Claim 2, wherein the third promoter comprises a CREB-binding element or a 19bp repeat from a hCMV-MIE enhancer.
- 20 17. The vector system of Claim 2, wherein the third promoter comprises a TATAA transcription initiation signal.
18. The vector system of Claim 2, wherein the third promoter is a hCMV-MIE promoter having a TATAA box region.
19. The vector system of Claim 2, wherein the third cistron is associated with a
25 ubiquitous chromatin opening element, an insulator, or a barrier element.